(FILE 'HOME' ENTERED AT 16:57:27 ON 09 NOV 2002)

FILE 'STNGUIDE' ENTERED AT 16:57:40 ON 09 NOV 2002 SET LINE 250

SET DETAIL OFF

FILE 'HOME' ENTERED AT 16:57:49 ON 09 NOV 2002

SET LINE LOGIN SET DETAIL LOGIN

FILE 'BIOSIS, AGRICOLA, ALUMINIUM, ANABSTR, AQUIRE, BABS, BIOCOMMERCE, BIOTECHNO, CABA, CAOLD, CAPLUS, CBNB, CEABA-VTB, CEN, CERAB, CIN, COMPENDEX, CONFSCI, COPPERLIT, CORROSION, DKILIT, ENCOMPLIT, ENCOMPLIT2, FEDRIP, GENBANK, INSPEC, INSPHYS, INVESTEXT, ...' ENTERED AT 16:59:10 ON 09 NOV 2002

22534 SEA ADHESIONS

1. . .

1

L1

L2

L3

L4

2471 SEA L1 AND PREVENTION

29 SEA L2 AND GLUCOSE

1 SEA L3 AND DEGREE OF POLYMERIZATION

D L4 1

D L4 1 ALL

D L3 TI

D L3 1-29, TI

D L3 1, 2, 5, 6, 9-11, 14, 17, 20, 26-28, STD, AB

D L3 1, 2, 5, 6, 9-11, 14, 17, 20, 26-28 ALL

FILE HOME

FILE STNGUIDE

FILE CONTAINS CURRENT INFORMATION.

LAST RELOADED: Nov 1, 2002 (20021101/UP).

FILE BIOSIS

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 7 November 2002 (20021107/ED)

FILE AGRICOLA

FILE COVERS 1970 TO 9 Nov 2002 (20021109/ED)

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FILE ALUMINIUM

FILE LAST UPDATED: 6 NOV 2002 <20021106/UP>

FILE COVERS 1968 TO DATE.

FILE ANABSTR

FILE LAST UPDATED: 4 NOV 2002 <20021104/UP>

FILE COVERS 1980 TO DATE.

FILE AQUIRE

FILE COVERS 1915 TO 18 Aug 2002 (20020818/ED)

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FILE BABS

FILE LAST UPDATED: 29 AUG 2002 <20020829/UP>

FILE COVERS 1980 TO DATE.

FILE BIOCOMMERCE

FILE LAST UPDATED: 5 NOV 2002 <20021105/UP>

FILE BIOTECHNO

FILE LAST UPDATED: 7 NOV 2002 <20021107/UP>

FILE COVERS 1980 TO DATE.

>>> SIMULTANEOUS LEFT AND RIGHT TRUNCATION AVAILABLE IN /CT AND BASIC INDEX <<<

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FILE COVERS 1973 TO 8 Nov 2002 (20021108/ED)

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FILE CAOLD

FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

This file supports REG1stRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

FILE CAPLUS

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FILE COVERS 1907 - 9 Nov 2002 VOL 137 ISS 20 FILE LAST UPDATED: 7 Nov 2002 (20021107/ED)

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FILE CBNB

FILE LAST UPDATED: 8 NOV 2002 <20021108/UP>
FILE COVERS 1984 TO DATE.

FILE CEABA-VTB

FILE LAST UPDATED: 7 NOV 2002 <20021107/UP>
FILE COVERS 1966 TO DATE

FILE CEN

FILE COVERS 1991 TO 12 Jan 2001 (20010112/ED).

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FILE COVERS 1976 TO 23 MAY 1997 (970523/ED)

THIS FILE IS CURRENTLY NOT BEING UPDATED.

FILE CIN

FILE COVERS 1974 - 8 NOV 2002 (20021108/ED) VOL 31 ISS 46

FILE COMPENDEX

FILE LAST UPDATED: 6 NOV 2002

<20021106/UP>

FILE COVERS 1970 TO DATE.

FILE CONFSCI

FILE COVERS 1973 TO 4 Sep 2002 (20020904/ED)

FILE COPPERLIT

FILE LAST UPDATED: 26 SEP 2002 <20020926/UP>

FILE COVERS 1965 TO DATE

>>> Simultaneous left and right truncation available in the Basic Index <<<

FILE CORROSION

FILE LAST UPDATED: 25 SEP 2002 <20020925/UP>

FILE COVERS 1980 TO DATE.

FILE DKILIT

FILE LAST UPDATED: 29 OCT 2002 <20021029/UP>

FILE COVERS 1973 TO DATE.

FILE ENCOMPLIT

FILE COVERS 1964 TO 6 Nov 2002 (20021106/ED)

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FILE LAST UPDATED: 4 NOV 2002 <20021104/UP>

FILE COVERS 1969 TO DATE.

FILE INSPHYS

FILE LAST UPDATED: 23 DEC 94 <941223/UP>

FILE COVERS 1979 - 1994

>>> INSPHYS THESAURUS AVAILABLE IN FIELD /CT <<<

FILE INVESTEXT

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FILE KOSMET

FILE LAST UPDATED: 4 NOV 2002 <20021104/UP>

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FILE LAST UPDATED: 2 NOV 2002 <20021102/UP>
FILE COVERS 1964 TO DATE.

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FILE PAPERCHEM2

FILE COVERS 1967 TO 6 Nov 2002 (20021106/ED)

FILE PASCAL

FILE LAST UPDATED: 6 NOV 2002 <20021106/UP>
FILE COVERS 1984 TO DATE.

FILE PROMT

FILE COVERS 1978 TO 8 NOV 2002 (20021108/ED)

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FILE RAPRA

FILE LAST UPDATED: 8 NOV 2002 <20021108/UP>
FILE COVERS 1972 TO DATE

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FILE RUSSCI

FILE LAST UPDATED: 07 OCT 2002 <20021007/UP>

FILE SCISEARCH

FILE COVERS 1974 TO 8 Nov 2002 (20021108/ED)

FILE TULSA

FILE COVERS 1965 TO 6 NOV 2002 (20021106/ED)

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FILE TULSA2

FILE COVERS 1965 TO 6 NOV 2002 (20021106/ED)

FILE USAN

FILE COVERS 1953 THROUGH JULY 2002

USAN was reloaded July 28, 2002. Answer sets saved prior to this date are no longer be valid. Saved queries and L-number sessions have not been impacted.

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FILE WELDASEARCH

FILE LAST UPDATED: 5 NOV 2002 <

<20021105/UP>

FILE COVERS 1967 TO DATE

>>> SIMULTANEOUS LEFT AND RIGHT TRUNCATION AVAILABLE IN THE BASIC INDEX <<<

FILE WSCA

FILE LAST UPDATED: 25 OCT 2002

<20021025/UP>

FILE COVERS 1976 TO DATE

- ANSWER 1 OF 29 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L3Development of a novel glucose polymer solution (icodextrin) for adhesion prevention: Pre-clinical studies. ANSWER 2 OF 29 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. T.3 ΤI Use of fibrinolytic agents in the prevention of postoperative adhesion formation. L3 ANSWER 3 OF 29 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. Mitochondrial dysfunction and cytosekeletal disruption during chemical TI hypoxia to cultured rat hepatic sinusoidal endothelial cells: The pH paradox and cytoprotection by glucose, acidotic pH, and glycine. L3 ANSWER 4 OF 29 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. EFFECT OF CORTICO STEROIDS ON BLOOD PICTURE IN POST PARTURIENT BUFFALO. TIL3 ANSWER 5 OF 29 BIOTECHNO COPYRIGHT 2002 Elsevier Science B.V. ΤI Development of a novel glucose polymer solution (icodextrin) for adhesion prevention: Pre-clinical studies ANSWER 6 OF 29 BIOTECHNO COPYRIGHT 2002 Elsevier Science B.V. 1.3 Use of fibrinolytic agents in the prevention of postoperative TIadhesion formation ANSWER 7 OF 29 BIOTECHNO COPYRIGHT 2002 Elsevier Science B.V. L3TI Mitochondrial dysfunction and cytoskeletal disruption during chemical hypoxia to cultured rat hepatic sinusoidal endothelial cells: The pH paradox and cytoprotection by glucose, acidotic pH, and glycine
 - L3 ANSWER 8 OF 29 BIOTECHNO COPYRIGHT 2002 Elsevier Science B.V. Adjuvants in tubal surgery
 - L3 ANSWER 9 OF 29 CAOLD COPYRIGHT 2002 ACS
 - TI Intraperitoneal use of hypertonic glucose solutionprevention of adhesions
 - L3 ANSWER 10 OF 29 CAPLUS COPYRIGHT 2002 ACS
 - TI A randomized, controlled pilot study of the safety and efficacy of 4% icodextrin solution in the reduction of adhesions following laparoscopic gynecological surgery
 - L3 ANSWER 11 OF 29 CAPLUS COPYRIGHT 2002 ACS
 - TI Development of a novel **glucose** polymer solution (icodextrin) for adhesion **prevention**: Pre-clinical studies
 - L3 ANSWER 12 OF 29 CAPLUS COPYRIGHT 2002 ACS
 - TI Mitochondrial dysfunction and cytoskeletal disruption during chemical hypoxia to cultured rat hepatic sinusoidal endothelial cells: the pH paradox and cytoprotection by **glucose**, acidotic pH, and glycine
 - L3 ANSWER 13 OF 29 CAPLUS COPYRIGHT 2002 ACS
 - TI Crosslinkable polypeptide compositions and their use in delivery of biologically active agents to subjects
 - L3 ANSWER 14 OF 29 CAPLUS COPYRIGHT 2002 ACS
 - TI The intraperitoneal use of hypertonic **glucose** solution. An experimental study with reference to the **prevention** of adhesions
 - L3 ANSWER 15 OF 29 CEN COPYRIGHT 2002 ACS
 - TI Tissue Engineering
 The groundwork for developing biological substitutes for damaged tissue is being prepared by a new, rapidly evolving interdisciplinary field that

draws on the expertise of chemical engineers

- L3 ANSWER 16 OF 29 INVESTEXT COPYRIGHT 2002 TFS
- TI Biotechnology Industry Product Chart Industry Report
- L3 ANSWER 17 OF 29 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED.
- TIEN Use of fibrinolytic agents in the **prevention** of postoperative adhesion formation
- L3 ANSWER 18 OF 29 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED.
- TIEN Development of a novel **glucose** polymer solution (icodextrin) for adhesion **prevention**: pre-clinical studies
- L3 ANSWER 19 OF 29 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED.
- TIEN Mitochondrial dysfunction and cytoskeletal disruption during chemical hypoxia to cultured rat hepatic sinusoidal endothelial cells : The pH paradox and cytoprotection by **glucose**, acidotic pH, and glycine
- L3 ANSWER 20 OF 29 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED.
- TIEN Preventing recurrent postoperative adhesions: an experimental study in rats
- L3 ANSWER 21 OF 29 PROMT COPYRIGHT 2002 Gale Group
- TI Competition heats up in implanted devices. (various medical company products) (Statistical Data Included)
- L3 ANSWER 22 OF 29 PROMT COPYRIGHT 2002 Gale Group
- TI Cutting--edge Carbohydrates.
- L3 ANSWER 23 OF 29 PROMT COPYRIGHT 2002 Gale Group
- TI Best PIPELINES.
- L3 ANSWER 24 OF 29 PROMT COPYRIGHT 2002 Gale Group
- TI Never richer
- L3 ANSWER 25 OF 29 PROMT COPYRIGHT 2002 Gale Group
- TI Pharmacy update: Focus on diabetes
- L3 ANSWER 26 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- TI A randomized, controlled pilot study of the safety and efficacy of 4(icodextrin solution in the reduction of adhesions following laparoscopic gynaecological surgery
- L3 ANSWER 27 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- TI Development of a novel **glucose** polymer solution (icodextrin) for adhesion **prevention**: pre-clinical studies
- L3 ANSWER 28 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- TI Use of fibrinolytic agents in the **prevention** of postoperative adhesion formation
- L3 ANSWER 29 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- TI Mitochondrial dysfunction and cytoskeletal disruption during chemical hypoxia to cultured rat hepatic sinusoidal endothelial cells: The pH paradox and cytoprotection by **glucose**, acidotic pH, and glycine

L3 ANSWER 11 OF 29 CAPLUS COPYRIGHT 2002 ACS

AN 2000:617408 CAPLUS

DN 134:120900

TI Development of a novel **glucose** polymer solution (icodextrin) for adhesion **prevention**: Pre-clinical studies

AU Verco, Shelagh J. S.; Peers, Elizabeth M.; Brown, Colin B.; Rodgers, Kathleen E.; Roda, Norma; diZerega, Gere

CS ML Laboratories PLC, Leicestershire, LE8 4FA, UK

Human Reproduction (2000), 15(8), 1764-1772 CODEN: HUREEE; ISSN: 0268-1161

PB Oxford University Press

DT Journal

SO

LA English

Intra-abdominal adhesion formation causes significant post-operative AB morbidity. Controlled studies using animal models have been carried out to assess the tolerability and preventive efficacy of icodextrin solution (a biodegradable, biocompatible, glucose polymer). Reduction of adhesion formation was first evaluated in a rabbit double uterine horn model, applying 10-75 mL of 7.5 and 20%, or 50 mL of 2.5-20% icodextrin solution post-operatively. Significant increases in adhesion free sites (P < 0.005) were observed with vols. \geq 25 mL, and at concns. \geq 4%. Efficacy of 50 mL 4 and 20% icodextrin was then evaluated both during and after surgery, demonstrating significant redns. in adhesion formation (P < 0.002). In one study, intraplus post-operative use of 4% icodextrin produced the greatest reduction of non-surgical site adhesions; in others, the post-operative effect was predominant. Post-surgical administration of 50 mL 4% icodextrin in a rabbit sidewall model also resulted in more adhesion-free animals, and a significant reduction (P < 0.001) in areas of adhesion formation and reformation. In a rat infection potentiation model, 4% icodextrin produced no difference in mortality, abscess formation or overall abscess score. These data suggest that 4% icodextrin offers a well-tolerated and effective means of reducing post-surgical adhesion formation.

RE.CNT 52 THERE ARE 52 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 14 OF 29 CAPLUS COPYRIGHT 2002 ACS

AN 1940:48614 CAPLUS

DN 34:48614

OREF 34:7438f-g

TI The intraperitoneal use of hypertonic **glucose** solution. An experimental study with reference to the **prevention** of adhesions

AU Totten, H. P.

SO Surgery (1940), 8, 456-63

DT Journal

LA Unavailable

AB Hypertonic **glucose** in normal salt solution is entirely innocuous when given intraperitoneally to rabbits. It is completely absorbed within 24 hrs. It prevents the formation and re-formation of exptl. **adhesions** and produces a certain degree of nonspecific immunity in the peritoneum. When gross peritoneal contamination is present, hypertonic **glucose** hastens the spread of infection.

=> d L3 1, 2, 5, 6, 9-11, 14, 17, 20, 26-28 all L3 ANSWER 1 OF 29 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. AN 2000:415079 BIOSIS DN PREV200000415079 ΤI Development of a novel qlucose polymer solution (icodextrin) for adhesion prevention: Pre-clinical studies. ΑU Verco, Shelagh J. S. (1); Peers, Elizabeth M.; Brown, Colin B.; Rodgers, Kathleen E.; Roda, Norma; DiZerega, Gere CS (1) ML Laboratories PLC, Blaby, Leicestershire, LE8 4FA UK SO Human Reproduction (Oxford), (August, 2000) Vol. 15, No. 8, pp. 1764-1772. print. ISSN: 0268-1161. DTArticle English LA SL English AΒ Intra-abdominal adhesion formation causes significant post-operative morbidity. Controlled studies using animal models have been carried out to assess the tolerability and preventive efficacy of icodextrin solution (a biodegradable, biocompatible, glucose polymer). Reduction of adhesion formation was first evaluated in a rabbit double uterine horn model, applying 10-75 ml of 7.5 and 20%, or 50 ml of 2.5-20% icodextrin solution post-operatively. Significant increases in adhesion free sites (P < 0.005) were observed with volumes gtoreq 25 ml, and at concentrations gtoreq 4%. Efficacy of 50 ml 4 and 20% icodextrin was then evaluated both during and after surgery, demonstrating significant reductions in adhesion formation (P < 0.002). In one study, intraplus post-operative use of 4% icodextrin produced the greatest reduction of non-surgical site adhesions; in others, the post-operative effect was predominant. Post-surgical administration of 50 ml 4% icodextrin in a rabbit sidewall model also resulted in more adhesion-free animals, and a significant reduction (P < 0.001) in areas of adhesion formation and reformation. In a rat infection potentiation model, 4% icodextrin produced no difference in mortality, abscess formation or overall abscess score. These data suggest that 4% icodextrin offers a well-tolerated and effective means of reducing post-surgical adhesion formation. CC Biophysics - Biocybernetics *10515 Mathematical Biology and Statistical Methods *04500 Reproductive System - Physiology and Biochemistry *16504 Developmental Biology - Embryology - General and Descriptive *25502 BC Leporidae 86040 Hominidae 86215 IT Major Concepts Models and Simulations (Computational Biology); Reproduction IT Chemicals & Biochemicals icodextrin: development, glucose polymer solution IT Miscellaneous Descriptors adhesion prevention; double uterine horn model; rabbit sidewall model of adhesion formation and reformation ORGN Super Taxa Hominidae: Primates, Mammalia, Vertebrata, Chordata, Animalia; Leporidae: Lagomorpha, Mammalia, Vertebrata, Chordata, Animalia ORGN Organism Name human (Hominidae): patient; rabbit (Leporidae) ORGN Organism Superterms Animals; Chordates; Humans; Lagomorphs; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Primates; Vertebrates RN 9004-53-9 (ICODEXTRIN) ANSWER 2 OF 29 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L3 2000:411246 BIOSIS ANDN PREV200000411246 ΤI Use of fibrinolytic agents in the prevention of postoperative

adhesion formation. ΑU Hellebrekers, Bart W. J. (1); Trimbos-Kemper, Trudy C. M.; Trimbos, J. Baptist M. Z.; Emeis, Jef J.; Kooistra, Teake CS (1) Department of Gynecology, Leiden University Medical Center, 2300 RC, Leiden Netherlands Fertility and Sterility, (August, 2000) Vol. 74, No. 2, pp. 203-212. SO print. ISSN: 0015-0282. DTGeneral Review LA English $_{ t SL}$ English AΒ Objective: To review the events leading to the formation of adhesions, to describe the development of fibrinolytic agents, to review more than a century of research on the use of fibrinolytic agents in adhesion prevention, and to look at future aspects of adhesion prevention. Results: A better understanding of the pathogenesis of adhesion formation has resulted in the use of fibrinolytic agents in their prevention. Fibrinolytic agents promote fibrinolytic activity during the early period after peritoneal trauma during which an increased formation of fibrin is seen in combination with a deficiency of endogenous fibrinolytic activity. Initially, chemical attacks on fibrin (fibrolysin and hypertonic glucose), foreign digestive ferments (pepsin, trypsin, and papain), and stimulation of intraperitoneal leukocytosis (amniotic fluid) were used. Development of new thrombolytic agents was soon followed by experiments in animal adhesion models and clinical studies to examine their antiadhesion properties. Plasmin preparations (plasmin, actase, and fibrinolysin) and plasmin activators (streptokinase, urokinase, and tissue-type plasminogen activator) were found to be efficacious in preventing adhesion formation in the greater part of reviewed animal and clinical studies. Conclusion(s): From the current literature, it can be concluded that postoperative intraperitoneal administration of thrombolytic agents can significantly decrease adhesion formation. Given the large number of experimental studies in animals, future studies should focus on the clinical use of fibrinolytic agents in the prevention of postsurgical adhesion formation. Digestive System - Pathology *14006 Biochemical Studies - Proteins, Peptides and Amino Acids *10064 Enzymes - General and Comparative Studies; Coenzymes *10802 Pathology, General and Miscellaneous - Therapy *12512 Blood, Blood-Forming Organs and Body Fluids - Blood, Lymphatic and Reticuloendothelial Pathologies *15006 Reproductive System - Pathology *16506 Pharmacology - General *22002 Pharmacology - Clinical Pharmacology *22005 Pharmacology - Blood and Hematopoietic Agents *22008 BC Hominidae 86215 IT Major Concepts Gynecology (Human Medicine, Medical Sciences); Hematology (Human Medicine, Medical Sciences); Pharmacology IT Diseases abdominal pelvic adhesion: digestive system disease, reproductive system disease/female; peritoneal trauma: injury IT Chemicals & Biochemicals actase: thrombolytic; fibrin; fibrinolysin: plasmin preparation, thrombolytic - drug; fibrinolytic agents: thrombolytic; fibrolysin; hypertonic glucose; papain; pepsin; plasmin: plasmin preparation, thrombolytic - drug; streptokinase: plasmin activators, thrombolytic - drug; thrombolytic agents: thrombolytic; tissue-type plasminogen activator: plasmin activator, thrombolytic - drug; trypsin; urokinase: plasmin activator, thrombolytic - drug IT Methods & Equipment intraperitoneal leukocytosis stimulation: therapeutic method IT Miscellaneous Descriptors

fibrinolysis; postoperative adhesion formation prevention ORGN Super Taxa Hominidae: Primates, Mammalia, Vertebrata, Chordata, Animalia ORGN Organism Name human (Hominidae): female, patient ORGN Organism Superterms Animals; Chordates; Humans; Mammals; Primates; Vertebrates 9001-90-5 (ACTASE) RN9001-90-5 (FIBRINOLYSIN) 8013-20-5 (FIBROLYSIN) 9001-73-4 (PAPAIN) 9001-75-6 (PEPSIN) 9001-90-5 (PLASMIN) 9002-01-1 (STREPTOKINASE) 139639-23-9 (TISSUE-TYPE PLASMINOGEN ACTIVATOR) 9002-07-7 (TRYPSIN) 9039-53-6 (UROKINASE) ANSWER 5 OF 29 L_3 BIOTECHNO COPYRIGHT 2002 Elsevier Science B.V. AN 2000:30636066 BIOTECHNO Development of a novel glucose polymer solution (icodextrin) TI for adhesion prevention: Pre-clinical studies ΑU Verco S.J.S.; Peers E.M.; Brown C.B.; Rodgers K.E.; Roda N.; DiZerega G. CS S.J.S. Verco, ML Laboratories PLC, Blaby, Leicestershire LE8 4FA, United Kingdom. E-mail: sjsverco@aol.com SO Human Reproduction, (2000), 15/8 (1764-1772), 52 reference(s) CODEN: HUREEE ISSN: 0268-1161 DT Journal; Article CY United Kingdom LΑ English SLEnglish AB Intra-abdominal adhesion formation causes significant post-operative morbidity. Controlled studies using animal models have been carried out to assess the tolerability and preventive efficacy of icodextrin solution (a biodegradable, biocompatible, glucose polymer). Reduction of adhesion formation was first evaluated in a rabbit double uterine horn model, applying 10-75 ml of 7.5 and 20%, or 50 ml of 2.5-20% icodextrin solution post-operatively. Significant increases in adhesion free sites (P < 0.005) were observed with volumes >=25 ml, and at concentrations >=4%. Efficacy of 50 ml 4 and 20% icodextrin was then evaluated both during and after surgery, demonstrating significant reductions in adhesion formation (P < 0.002). In one study, intraplus post-operative use of 4% icodextrin produced the greatest reduction of non-surgical site adhesions; in others, the post-operative effect was predominant. Post-surgical administration of 50 ml 4% icodextrin in a rabbit sidewall model also resulted in more adhesion-free animals, and a significant reduction (P < 0.001) in areas of adhesion formation and reformation. In a rat infection potentiation model, 4% icodextrin produced no difference in mortality, abscess formation or overall abscess score. These data suggest that 4% icodextrin offers a well-tolerated and effective means of reducing post-surgical adhesion formation. CT*icodextrin; *peritoneum adhesion; controlled study; animal model; nonhuman; female; uterus synechia; postoperative complication; drug efficacy; drug tolerability; rabbit; uterus; bacterial peritonitis; article CO Drug Manufacturer: ML, United Kingdom L3ANSWER 6 OF 29 BIOTECHNO COPYRIGHT 2002 Elsevier Science B.V. AN 2000:30604178 **BIOTECHNO** TI Use of fibrinolytic agents in the prevention of postoperative adhesion formation ΑU Hellebrekers B.W.J.; Trimbos-Kemper T.C.M.; Trimbos J.B.M.Z.; Emeis J.J.; Kooistra T.

CS Dr. B.W.J. Hellebrekers, Department of Gynecology, Leiden University Medical Center, P.O. Box 9600, 2300 RC Leiden, Netherlands. E-mail: bwjh@xs4all.nl Fertility and Sterility, (2000), 74/2 (203-212), 104 reference(s) SO CODEN: FESTAS ISSN: 0015-0282 PUI S0015028200006567 DTJournal; General Review United States CY LAEnglish SL English Objective: To review the events leading to the formation of AB adhesions, to describe the development of fibrinolytic agents, to review more than a century of research on the use of fibrinolytic agents in adhesion prevention, and to look at future aspects of adhesion prevention. Results: A better understanding of the pathogenesis of adhesion formation has resulted in the use of fibrinolytic agents in their prevention. Fibrinolytic agents promote fibrinolytic activity during the early period after peritoneal trauma during which an increased formation of fibrin is seen in combination with a deficiency of endogenous fibrinolytic activity. Initially, chemical attacks on fibrin (fibrolysin and hypertonic glucose), foreign digestive ferments (pepsin, trypsin, and papain), and stimulation of intraperitoneal leukocytosis (amniotic fluid) were used. Development of new thrombolytic agents was soon followed by experiments in animal adhesion models and clinical studies to examine their antiadhesion properties. Plasmin preparations (plasmin, actase, and fibrinolysin) and plasmin activators (streptokinase, urokinase, and tissue-type plasminogen activator) were found to be efficacious in preventing adhesion formation in the greater part of reviewed animal and clinical studies. Conclusion(s): From the current literature, it can be concluded that postoperative intraperitoneal administration of thrombolytic agents can significantly decrease adhesion formation. Given the large number of experimental studies in animals, future studies should focus on the clinical use of fibrinolytic agents in the prevention of postsurgical adhesion formation. Copyright (C) 2000 American Society for Reproductive Medicine. CT*fibrinolytic agent; *fibrinolysis; *postoperative complication; *peritoneum adhesion; *adhesion; streptokinase; plasmin; varidase; plasminogen activator; reteplase; pepsin A; papain; alteplase; trypsin; urokinase; fibrolysin; unclassified drug; peritoneum; human; nonhuman; review; priority journal RN(streptokinase) 9002-01-1; (plasmin) 9001-90-5, 9004-09-5; (varidase) 8048-16-6; (plasminogen activator) 9039-53-6; (reteplase) 133652-38-7; (pepsin A) 9001-75-6; (papain) 9001-73-4; (alteplase) 105857-23-6; (trypsin) 9002-07-7; (urokinase) 139639-24-0 ANSWER 9 OF 29 CAOLD COPYRIGHT 2002 ACS L3 AN CA34:7438g CAOLD Intraperitoneal use of hypertonic glucose solution-ŤΤ prevention of adhesions ΑU Totten, H. P. ANSWER 10 OF 29 CAPLUS COPYRIGHT 2002 ACS L3AN 2002:369128 CAPLUS DN 137:57517 TΙ A randomized, controlled pilot study of the safety and efficacy of 4% icodextrin solution in the reduction of adhesions following laparoscopic gynecological surgery AU DiZerega, G. S.; Verco, S. J. S.; Young, P.; Kettel, M.; Kobak, W.; Martin, D.; Sanfilippo, J.; Peers, E. M.; Scrimgeour, A.; Brown, C. B.

University of Southern California Keck School of Medicine, Los Angeles,

SO Human Reproduction (2002), 17(4), 1031-1038 CODEN: HUREEE; ISSN: 0268-1161

CS

PΒ Oxford University Press \mathbf{DT} Journal LA English CC 1-12 (Pharmacology) AΒ BACKGROUND: Adhesion-related readmissions are frequent sequelae to gynaecol. surgery. Attempts to prevent adhesions by separating healing peritoneal surfaces include site-specific barriers and hydroflotation by instilled solns. Rapid absorption limits the effectiveness of solns. such as Ringer's lactated saline (RLS). pilot study assessed the safety, tolerability and preliminary effectiveness of a non-viscous, iso-osmolar solution of 4% icodextrin, an α -1,4 **glucose** polymer with prolonged i.p. residence, in reducing adhesions after laparoscopic gynaecol. surgery. METHODS: Women aged ≥18 yr, requiring laparoscopic adnexal surgery (n = 62), were entered into a randomized, open-label, assessor-blinded, multicenter study to compare 4% icodextrin with RLS. Treatments were coded in blocks of four with equal randomization to each group, and pre-allocated to consecutively numbered patients. At least 100 mL per 30 min was used for intra-operative lavage, with 1 l instilled post-operatively. Per protocol anal. included all eligible patients (n =53); reformation anal. required one or more baseline adhesion (n = 42). Incidence, extent and severity of post-operative adhesions were assessed at second-look laparoscopy after 6-12 wk. Procedures were video-taped for third party, blinded assessment. RESULTS: Safety and tolerability (laboratory variables, adverse events, clin. follow-up) were good with no difference between treatments. A shift anal. of incidence-ranked adhesions (n = 53) showed apparent improvements in more patients with icodextrin than RLS (37 vs. 15%; not significant). Adhesion score reduction (n = 42) was more frequent in icodextrin- than RLS-treated patients: incidence (52 vs. 32%), extent (52 vs. 47%), and severity (65 vs. 37%). Despite greater baseline adhesions, median reformation was less after icodextrin (24%) than RLS (60%). The pilot study group sizes were not powered for statistical significance. CONCLUSIONS: In this preliminary study, 4% icodextrin lavage plus instillation was well tolerated and reduced adhesion formation and reformation following laparoscopic gynaecol. surgery. A Phase III pivotal study is currently in progress. ST icodextrin soln peritoneal adhesion redn prevention laparoscopy gynecol surgery IT Named reagents and solutions RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (Ringer's lactate; icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) IT Peritoneum (adhesion; icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) ΙT Peritoneum (cavity, instillation of icodextrin solution; icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) ITHuman (icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) TТ Abdomen Surgery (laparoscopy, gynecol.; icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) IT Adhesion, biological

(peritoneal; icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) \mathbf{T} 337376-15-5, Adept RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (icodextrin solution (Adept) vs. Ringers lactate: safety and efficacy in reduction of patient adhesions following laparoscopic gynaecol. surgery) RE.CNT 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) American Fertility Society; Fertil Steril 1988, V49, P944 (2) Damario, M; J Gynecol Technol 1995, V1, P77 (3) Davies, D; Perit Dial Int 1994, V14, PS45 (4) Dawnay, A; Perit Dial Int 1997, V17, P52 MEDLINE (5) diZerega, G; Eur J Surg 1997, V163 (Suppl 577), P10 (6) diZerega, G; Fertil Steril 1994, V61, P219 MEDLINE (7) diZerega, G; Infertil Reprod Med Clinics North Am 1994, V5, P463 (8) diZerega, G; Pelvic Surgery Adhesion Formation and Prevention 1997, P188 (9) Diamond, M; Fertil Steril 1996, V66, P904 MEDLINE (10) Diamond, M; Fertil Steril 1998, V69, P1067 MEDLINE (11) Dobbie, J; Eur Pharm Contractor 1999, P44 (12) Ellis, H; Lancet 1999, V353, P1476 MEDLINE (13) Gauwerky, J; Biol Res Pregnancy 1986, V7, P93 MEDLINE (14) Gilbert, J; Perit Dial Int 1999, V19(Suppl 1), PS78 (15) Gilbert, J; Perit Dial Int 1999, V19(Suppl 1), PS79 (16) Hart, R; Gynaecol Endosc 1996, V5, P287 (17) Holmdahl, L; Eur J Surg 1997, V163 (Suppl 577), P56 (18) Holmdahl, L; Surgery 1998, V123, P539 MEDLINE (19) Hosie, K; Drug Delivery 2001, V8, P9 CAPLUS (20) Interceed (TC7) Adhesion Barrier Study Group; Fertil Steril 1989, V51, P933 (21) Johns, D; Fertil Steril 1999, V72 (Suppl 1), PS57 (22) Johns, D; Peritoneal Surgery 1999, P351 (23) Kerr, D; Br J Cancer 1996, V74, P2032 CAPLUS (24) Lower, A; Br J Obstet Gynaecol 2000, V107, P855 MEDLINE (25) McArdle, C; Br J Cancer 1994, V70, P762 MEDLINE (26) Mistry, C; Kidney Int 1994, V46, P496 MEDLINE (27) Peers, E; Perit Dial Int 1997, V17, P22 MEDLINE (28) Posthuma, N; Nephrol Dial Transplant 1997, V12, P550 CAPLUS (29) Shear, L; N Engl J Med 1965, V272, P123 MEDLINE (30) Sites, C; J Ultrasound Med 1997, V16, P195 MEDLINE (31) Stout, A; Am J Obstet Gynecol 1991, V146, P73 (32) Strickler, B; J Am Coll Surg 1994, V178, P617 (33) Surgical Membrane Study Group; Fertil Steril 1992, V57, P921 (34) Thornton, M; Hum Reprod 1998, V13, P1480 CAPLUS (35) Thornton, M; Peritoneal Adhesions 1997, P370 (36) Topley, N; Perit Dial Int 1994, V14, PS28 (37) Trimbos-Kemper, T; Fertil Steril 1985, V43, P396 (38) Verco, S; Eur Pharm Contractor 2000, P74 (39) Verco, S; Hum Reprod 2000, V15, P1764 CAPLUS (40) Wiseman, D; Fertil Steril 1998, V70, P702 MEDLINE L3 ANSWER 11 OF 29 CAPLUS COPYRIGHT 2002 ACS AN2000:617408 CAPLUS DN 134:120900 TΤ Development of a novel glucose polymer solution (icodextrin) for adhesion prevention: Pre-clinical studies ΑU Verco, Shelagh J. S.; Peers, Elizabeth M.; Brown, Colin B.; Rodgers, Kathleen E.; Roda, Norma; diZerega, Gere CS ML Laboratories PLC, Leicestershire, LE8 4FA, UK SO Human Reproduction (2000), 15(8), 1764-1772 CODEN: HUREEE; ISSN: 0268-1161 PB Oxford University Press

DT Journal LA English CC 63-8 (Pharmaceuticals) AB Intra-abdominal adhesion formation causes significant post-operative Controlled studies using animal models have been carried out to assess the tolerability and preventive efficacy of icodextrin solution (a biodegradable, biocompatible, glucose polymer). Reduction of adhesion formation was first evaluated in a rabbit double uterine horn model, applying 10-75 mL of 7.5 and 20%, or 50 mL of 2.5-20% icodextrin solution post-operatively. Significant increases in adhesion free sites (P < 0.005) were observed with vols. \geq 25 mL, and at concns. \geq 4%. Efficacy of 50 mL 4 and 20% icodextrin was then evaluated both during and after surgery, demonstrating significant redns. in adhesion formation (P < In one study, intraplus post-operative use of 4% icodextrin produced the greatest reduction of non-surgical site adhesions; in others, the post-operative effect was predominant. Post-surgical administration of 50 mL 4% icodextrin in a rabbit sidewall model also resulted in more adhesion-free animals, and a significant reduction (P < 0.001) in areas of adhesion formation and reformation. In a rat infection potentiation model, 4% icodextrin produced no difference in mortality, abscess formation or overall abscess score. These data suggest that 4% icodextrin offers a well-tolerated and effective means of reducing post-surgical adhesion formation. ST icodextrin soln peritoneum adhesion Adhesion, biological ΙT Peritoneum (pre-clin. studies of biodegradable and biocompatible icodextrin solns. for adhesion prevention) TT Drug delivery systems (solns.; pre-clin. studies of biodegradable and biocompatible icodextrin solns. for adhesion prevention) TΥ 9004-53-9, Icodextrin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (pre-clin. studies of biodegradable and biocompatible icodextrin solns. for adhesion **prevention**) THERE ARE 52 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT (1) Azziz, R; Surg Gynecol Obstet 1993, V177, P135 MEDLINE (2) Beck, D; Dis Colon Rectum 1999, V42, P241 MEDLINE (3) Bhatia, D; Am Surg 1997, V63, P775 MEDLINE (4) Bronson, R; Fertil Steril 1977, V28, P613 (5) Davies, D; Perit Dial Int 1994, V14, PS45 (6) diZerega, G; Fertil Steril 1994, V61, P219 MEDLINE (7) diZerega, G; Infertil Reprod Med Clinics N Am 1994, V5, P463 (8) diZerega, G; The Peritoneum 1992, P2 (9) diZerega, G; The Peritoneum 1992, P325 (10) diZerega, G; The Peritoneum 1992, P378 (11) Diamond, M; Fertil Steril 1988, V69, P1067 (12) Diamond, M; Fertil Steril 1996, V66, P904 MEDLINE (13) Ellis, H; Lancet 1999, V353, P1476 MEDLINE (14) Fayez, J; Am J Obstet Gynecol 1987, V157, P1184 MEDLINE (15) Franklin, R; Obstet Gynecol 1995, V86, P335 MEDLINE (16) Gauwerky, J; Biol Res Pregnancy Perinatol 1986, V7, P93 MEDLINE (17) Gilbert, J; Perit Dial Int 1999, V19, PS79 (18) Harris, E; Surgery 1995, V117, P663 MEDLINE (19) Hart, R; Gynaecol Endosc 1996, V5, P287 (20) Ho-dac-Pannekeet, M; Kidney Int 1996, V50, P979 MEDLINE (21) Holmdahl, L; Lancet 1999, V353, P1456 MEDLINE (22) Howard, F; Obstet Gynecol Surv 1993, V48, P357 MEDLINE (23) Interceed (TC7) Adhesion Barrier Study Group; Fertil Steril 1989, V51, P933 (24) Johns, D; Fertil Steril 1997, V68, P37 MEDLINE

(25) Keckstein, J; Hum Reprod 1996, V11, P579 MEDLINE (26) Kresch, A; Obstet Gynecol 1984, V64, P672 MEDLINE (27) Liberek, T; Nephron 1993, V65, P260 MEDLINE (28) Lindenberg, S; Eur Surg Res 1982, V14, P274 MEDLINE (29) Mais, V; Hum Reprod 1995, V10, P3133 MEDLINE (30) Miller, E; Arch Surg 1959, V78, P148 (31) Monk, B; Am J Obstet Gynecol 1994, V170, P1396 MEDLINE (32) Naether, O; Fertil Steril 1993, V60, P95 MEDLINE (33) Nishimura, K; Am J Med 1984, V77, P102 CAPLUS (34) Nordic Adhesion Prevention Study Group; Fertil Steril 1995, V63, P709 (35) Raftery, A; J Anat 1973, V115, P375 (36) Rodgers, K; Fertil Steril 1998, V70, P1131 MEDLINE (37) Rodgers, K; Fundam Appl Toxicol 1997, V36, P1 CAPLUS (38) Rodgers, K; Hum Reprod 1998, V12, P2443 (39) Rodgers, K; J Invest Surg 1996, V9, P388 (40) Rodgers, K; J Invest Surg 1997, V10, P31 MEDLINE (41) Shear, L; N Engl J Med 1965, V272, P123 MEDLINE (42) Sitter, T; Thromb Haemost 1999, V82, P1171 CAPLUS (43) Stout, A; Am J Obstet Gynecol 1991, V146, P73 (44) Strickler, B; J Am Coll Surg 1994, V178, P617 (45) Thomas, S; Am J Kidney Dis 1997, V29, P246 CAPLUS (46) Thornton, M; Hum Reprod 1998, V13, P1480 CAPLUS (47) Topley, N; Perit Dial Int 1994, V14, PS28 (48) Trimbos-Kemper, T; Fertil Steril 1989, V51, P1053 MEDLINE (49) Tulandi, T; Obstet Gynecol 1998, V92, P766 MEDLINE (50) Verco, S; Hum Reprod 1999, V14, P275 (51) Weinstein, W; Infect Immunol 1974, V10, P1250 MEDLINE (52) Wiseman, D; Fertil Steril 1998, V70, P702 MEDLINE L3 ANSWER 14 OF 29 CAPLUS COPYRIGHT 2002 ACS AN 1940:48614 CAPLUS DN 34:48614 OREF 34:7438f-q The intraperitoneal use of hypertonic glucose solution. An experimental study with reference to the prevention of adhesions ΑU Totten, H. P. Surgery (1940), 8, 456-63 SO DT Journal LA Unavailable CC 11H (Biological Chemistry: Pharmacology) AB Hypertonic glucose in normal salt solution is entirely innocuous when given intraperitoneally to rabbits. It is completely absorbed within 24 hrs. It prevents the formation and re-formation of exptl. adhesions and produces a certain degree of nonspecific immunity in the peritoneum. When gross peritoneal contamination is present, hypertonic glucose hastens the spread of infection. 1.3 ANSWER 17 OF 29 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED. AN 2000-0455327 PASCAL CP Copyright .COPYRGT. 2000 INIST-CNRS. All rights reserved. TIEN Use of fibrinolytic agents in the prevention of postoperative adhesion formation AU HELLEBREKERS B. W. J.; TRIMBOS-KEMPER T. C. M.; TRIMBOS J. B. M. Z.; EMEIS J. J.; KOOISTRA T. CS Leiden University Medical Center, Leiden, Netherlands; Gaubius Laboratory TNO-PG, Leiden, Netherlands so Fertility and sterility, (2000), 74(2), 203-212, 104 refs. ISSN: 0015-0282 CODEN: FESTAS DT Journal BLAnalytic CY United States LA English ΑV INIST-4120, 354000091007290010

- AΒ Objective: To review the events leading to the formation of adhesions, to describe the development of fibrinolytic agents, to review more than a century of research on the use of fibrinolytic agents in adhesion prevention, and to look at future aspects of adhesion prevention. Results: A better understanding of the pathogenesis of adhesion formation has resulted in the use of fibrinolytic agents in their prevention. Fibrinolytic agents promote fibrinolytic activity during the early period after peritoneal trauma during which an increased formation of fibrin is seen in combination with a deficiency of endogenous fibrinolytic activity. Initially, chemical attacks on fibrin (fibrolysin and hypertonic glucose). foreign digestive ferments (pepsin, trypsin, and papain), and stimulation of intraperitoneal leukocytosis (amniotic fluid) were used. Development of new thrombolytic agents was soon followed by experiments in animal adhesion models and clinical studies to examine their antiadhesion properties. Plasmin preparations (plasmin, actase, and fibrinolysin) and plasmin activators (streptokinase, urokinase, and tissue-type plasminogen activator) were found to be efficacious in preventing adhesion formation in the greater part of reviewed animal and clinical studies. Conclusion(s): From the current literature, it can be concluded that postoperative intraperitoneal administration of thrombolytic agents can significantly decrease adhesion formation. Given the large number of experimental studies in animals, future studies should focus on the clinical use of fibrinolytic agents in the prevention of postsurgical adhesion formation.
- CC 002B02G; Life sciences; Medical sciences; Pharmacology; Hematology CT Postoperative; Fibrinolytic; Pepsin A; Trypsin; Papain; Streptokinase; Plasmin; u-Plasminogen activator; t-Plasminogen activator; Review; Adhesion
- BT Aspartic endopeptidases; Peptidases; Hydrolases; Enzyme; Serine endopeptidases; Cysteine endopeptidases; Prevention; Fibrinolysis; Mechanism of action
- L3 ANSWER 20 OF 29 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED.

AN 1990-0183104 PASCAL

- TIEN Preventing recurrent postoperative adhesions: an experimental study in rats
- AU VERREET P. R.; FAKIR C.; OHMANN C.; ROEHER H. D.
- CS Heinrich-Heine-univ., dep. surgery, Duesseldorf 4000, Germany, Federal Republic of
- SO European Surgical Research, (1989), 21(5), 267-273, 20 refs. ISSN: 0014-312X CODEN: EUSRBM
- DT Journal
- BL Analytic
- CY Switzerland
- LA English
- AV CNRS-14606
- AB A peritoneal lavage model, cyclic intraperitoneal lavage (CIPL), and other adhesion preventing methods with and without fibrinolytic agents were compared to a control group without treatment in an animal study. The adhesion-preventing effect was evaluated at the site of a standardized peritoneal defect (free peritoneal grafting, P) and at the laparotomy wound (L) of 60 rats (12 escape) after surgical lysis of primary adhesions during relaparotomy
- CC 002B25G04; Life sciences; Medical sciences; Gastroenterology, Digestive system
- CT Adhesion; Peritoneum; Postoperative; **Prevention**; Washing; Peritoneal cavity; Ringer solution; **Glucose**; Fibrinolytic; Digestive diseases; Animal model; Animal; Rat
- BT Rodentia; Mammalia; Vertebrata
- L3 ANSWER 26 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- AN 2002:358572 SCISEARCH
- GA The Genuine Article (R) Number: 543UA

- TI A randomized, controlled pilot study of the safety and efficacy of 4(icodextrin solution in the reduction of adhesions following laparoscopic gynaecological surgery
- AU diZerega G S; Verco S J S (Reprint); Young P; Kettel M; Kobak W; Martin D; Sanfilippo J; Peers E M; Scrimgeour A; Brown C B
- CS ML Labs PLC, Blaby Hall, Church St, Leicester LE8 4FA, Leics, England (Reprint); ML Labs PLC, Leicester LE8 4FA, Leics, England; Univ So Calif, Keck Sch Med, Los Angeles, CA USA; IGO Med Grp, San Diego, CA USA; San Diego Fertil Ctr, San Diego, CA USA; Allegheny Gen Hosp, Pittsburgh, PA 15212 USA
- CYA England; USA
- SO HUMAN REPRODUCTION, (APR 2002) Vol. 17, No. 4, pp. 1031-1038.
 Publisher: OXFORD UNIV PRESS, GREAT CLARENDON ST, OXFORD OX2 6DP, ENGLAND.
 ISSN: 0268-1161.
- DT Article; Journal
- LA English

RE

- REC Reference Count: 40
- Background: Adhesion-related readmissions are frequent sequelae to AB gynaecological surgery. Attempts to prevent adhesions by separating healing peritoneal surfaces include site-specific barriers and hydroflotation by instilled solutions. Rapid absorption limits the effectiveness of solutions such as Ringer's lactated saline (RLS). This pilot study assessed the safety, tolerability and preliminary effectiveness of a non-viscous, iso-osmolar solution of 4% icodextrin, an alpha-1,4 glucose polymer with prolonged intraperitoneal residence, in reducing adhesions after laparoscopic gynaecological surgery. Methods: Women aged greater than or equal to18 years, requiring laparoscopic adnexal surgery (n=62), were entered into a randomized, open-label, assessor-blinded, multicentre study to compare 4% icodextrin with RLS. Treatments were coded in blocks of four with equal randomization to each group, and pre-allocated to consecutively numbered patients. At least 100 ml per 30 min was used for intra-operative lavage, with 1 l instilled post-operatively. Per protocol analysis included all eligible patients (n=53); reformation analysis required one or more baseline adhesion (n=42). Incidence, extent and severity of post-operative adhesions were assessed at second-look laparoscopy after 6-12 weeks. Procedures were video-taped for third party, blinded assessment. Results: Safety and tolerability (laboratory variables, adverse events, clinical follow-up) were good with no difference between treatments. A shift analysis of incidence-ranked adhesions (n=53) showed apparent improvements in more patients with icodextrin than RLS (37 versus 15%; not significant). Adhesion score reduction (n=42) was more frequent in icodextrin-than RLS-treated patients: incidence (52 versus 32%), extent (52 versus 47%), and severity (65 versus 37%). Despite greater baseline adhesions, median reformation was less after icodextrin (24%) than RLS (60%). The pilot study group sizes were not powered for statistical significance. Conclusions: In this preliminary study, 4% icodextrin lavage plus instillation was well tolerated and reduced adhesion formation and reformation following laparoscopic gynaecological surgery. A Phase III pivotal study is currently in progress.
- CC OBSTETRICS & GYNECOLOGY; REPRODUCTIVE BIOLOGY
- ST Author Keywords: **glucose** polymer; icodextrin; laparoscopic surgery; peritoneal **adhesions**; post-surgical **adhesions**
- STP KeyWords Plus (R): HOSPITAL READMISSIONS; PELVIC-SURGERY; PREVENTION; MULTICENTER; GLUCOSE; 5-FLUOROURACIL

| Referenced Author (RAU) | Year (RPY) | , | PG (RPG) | Referenced Work (RWK) |
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| DAWNAY A B S J | 1997 | 117 | 152 | PERITON DIALYSIS INT |
|---------------------|------|-----|------------|----------------------|
| DIAMOND M P | 1998 | 69 | 1067 | FERTIL STERIL |
| DIAMOND M P | 1996 | 66 | 904 | FERTIL STERIL |
| DIZEREGA G S | 1994 | 61 | 219 | FERTIL STERIL |
| DIZEREGA G S | 1994 | 5 | 463 | INFERTIL REPROD MED |
| DIZEREGA G S | 1997 | 163 | 10 | EUR J SURG S577 |
| DIZEREGA G S | 1997 | İ | 188 | PELVIC SURG ADHESION |
| DOBBIE J W | 1999 | İ | 44 | EUR PHARM CONTRA MAY |
| ELLIS H | 1999 | 353 | 1476 | LANCET |
| GAUWERKY J F H | 1986 | 7 | 93 | BIOL RES PREG PERIN |
| GILBERT J A | 1999 | 19 | S79 | PERITON DIAL INT S1 |
| GILBERT J A | 1999 | 19 | S78 | PERITON DIALYSIS INT |
| HART R | 1996 | 5 | 287 | GYNAECOL ENDOSC |
| HOLMDAHL L | 1997 | 163 | 56 | EUR J SURG S577 |
| HOLMDAHL L | 1998 | 123 | 539 | SURGERY |
| HOSIE K | 2001 | 8 | j 9 | DRUG DELIV |
| JOHNS D B | 1999 | 72 | S57 | FERTIL STERIL S1 |
| JOHNS D B | 1999 | ĺ | 351 | PERITONEAL SURG |
| KERR D J | 1996 | 74 | 2032 | BRIT J CANCER |
| LOWER A M | 2000 | 107 | 855 | BRIT J OBSTET GYNAEC |
| MCARDLE C S | 1994 | 70 | 762 | BRIT J CANCER |
| MISTRY C D | 1994 | 46 | 496 | KIDNEY INT |
| PEERS E | 1997 | 17 | 22 | PERITON DIALYSIS INT |
| POSTHUMA N | 1997 | 12 | 550 | NEPHROL DIAL TRANSPL |
| SHEAR L | 1965 | 272 | 123 | NEW ENGL J MED |
| SITES C K . | 1997 | 16 | 195 | J ULTRAS MED |
| STOUT A L | 1991 | 146 | 73 | AM J OBSTET GYNECOL |
| STRICKER B | 1994 | 178 | 617 | J AM COLL SURGEONS |
| THORNTON M H | 1998 | 13 | 1480 | HUM REPROD |
| THORNTON M H | 1997 | | 370 | PERITONEAL ADHESIONS |
| TOPLEY N | 1994 | 14 | S28 | PERITON DIALYSIS INT |
| TRIMBOSKEMPER T C M | 1985 | 43 | 396 | FERTIL STERIL |
| VERCO S J S | 2000 | 15 | 1764 | HUM REPROD |
| VERCO S J S | 2000 | | 74 | EUR PHARM CONTRA AUG |
| WISEMAN D M | 1998 | 70 | 702 | FERTIL STERIL |

- L3 ANSWER 27 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- AN 2000:623477 SCISEARCH
- GA The Genuine Article (R) Number: 342KM
- TI Development of a novel **glucose** polymer solution (icodextrin) for adhesion **prevention**: pre-clinical studies
- AU Verco S J S (Reprint); Peers E M; Brown C B; Rodgers K E; Roda N; diZerega G
- CS ML LABS PLC, BLABY LE8 4FA, LEICS, ENGLAND (Reprint); LIVINGSTON RES INST, LOS ANGELES, CA 90033
- CYA ENGLAND; USA
- SO HUMAN REPRODUCTION, (AUG 2000) Vol. 15, No. 8, pp. 1764-1772.
 Publisher: OXFORD UNIV PRESS, GREAT CLARENDON ST, OXFORD OX2 6DP, ENGLAND.
 ISSN: 0268-1161.
- DT Article; Journal
- FS LIFE; CLIN
- LA English
- REC Reference Count: 52
- AB Intra-abdominal adhesion formation causes significant post-operative morbidity, Controlled studies using animal models have been carried out to assess the tolerability and preventive efficacy of icodextrin solution (a biodegradable, biocompatible, glucose polymer). Reduction of adhesion formation was first evaluated in a rabbit double uterine horn model, applying 10-75 ml of 7.5 and 20%, or 50 ml of 2.5-20% icodextrin solution post-operatively. Significant increases in adhesion free sites (P < 0.005) were observed with volumes greater than or equal to 25 mi, and at concentrations greater than or equal to 4%. Efficacy of 50 mi 4 and 20% icodextrin was then evaluated both during and after surgery, demonstrating significant reductions in adhesion formation (P < 0.002), In one study,

intraplus post-operative use of 4% icodextrin produced the greatest reduction of non-surgical site adhesions; in others, the post-operative effect was predominant. Post-surgical administration of 50 mi 4% icodextrin in a rabbit sidewall model also resulted in more adhesion-free animals, and a significant reduction (P < 0.001) in areas of adhesion formation and reformation. In a rat infection potentiation model, 4% icodextrin produced no difference in mortality, abscess formation or overall abscess score. These data suggest that 4% icodextrin offers a well-tolerated and effective means of reducing post-surgical adhesion formation.

CC REPRODUCTIVE BIOLOGY; OBSTETRICS & GYNECOLOGY

ST Author Keywords: **glucose** polymer; icodextrin; peritoneal **adhesions**; preclinical

STP KeyWords Plus (R): INDUCED POSTOPERATIVE ADHESIONS;
PELVIC-SURGERY; REDUCTION; OBSTRUCTION; MYOMECTOMY; BARRIER

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| Deferenced Author | 1 37 | 1 7707 | l na | l m.c |
|---|------|-------------|---------------|----------------------|
| | | | PG | Referenced Work |
| (RAU) | | (RVL) | • | (RWK) |
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| *NORD ADH PREV STU | 1995 | 63 | 709 | FERTIL STERIL |
| AZZIZ R | 1993 | 177 | 135 | SURG GYNECOL OBSTET |
| BECK D E | 1999 | 42 | 241 | DIS COLON RECTUM |
| BHATIA D S | 1997 | 63 | 775 | AM SURGEON |
| BRONSON R A | 1977 | 28 | 613 | FERTIL STERIL |
| DAVIS D | 1994 | 14 | 4 | PUBLIC MONEY MANAGE |
| DIAMOND M | 1988 | 69 | 1067 | FERTIL STERIL |
| DIAMOND M P | 1996 | 66 | 904 | FERTIL STERIL |
| DIZEREGA G S | 1994 | 61 | 219 | FERTIL STERIL |
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| DIZEREGA G S | 1992 | ! [| 325 | PERITONEUM |
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| ELLIS H | ! | 252 | 376 1476 | ! |
| FAYEZ J A | ! | 353 157 | ! | LANCET |
| FRANCOIS M | ! | ! | 1184 | AM J OBSTET GYNECOL |
| | 1995 | 2 | 86 | ARCH PEDIATR |
| GAUWERKY J F H | ! | 7 | 93 | BIOL RES PREG PERIN |
| GILBERT J A | ! | 19 | S79 | PERITON DIALYSIS INT |
| HARRIS E S | ! | 117 | 663 | SURGERY |
| HART R | 1996 | 5 | 287 | GYNAECOL ENDOSC |
| HODACPANNEKEET M M | 1996 | 50 | 979 | KIDNEY INT |
| HOLMDAHL L | 1999 | 353 | 1456 | LANCET |
| HOWARD F M | 1993 | 48 | 357 | OBSTET GYNECOL SURV |
| JOHNS D B | 1997 | 68 | 37 | FERTIL STERIL |
| KECKSTEIN J | 1996 | 11 | 579 | HUM REPROD |
| KRESCH A J | 1984 | 64 | 672 | OBSTET GYNECOL |
| LIBEREK T | 1993 | 65 | 260 | NEPHRON |
| LINDENBERG S | 1982 | 14 | 274 | EUR SURG RES |
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| MONK B J | 1994 | 170 | 1396 | AM J OBSTET GYNECOL |
| NAETHER O G J | 1993 | 60 | 95 | FERTIL STERIL |
| NISHIMURA K | 1984 | 77 | 102 | AM J MED |
| RAFTERY A T | 1973 | 115 | 375 | J ANAT |
| RODGERS K | 1997 | 36 | 1 | FUND APPL TOXICOL |
| RODGERS K E | 1998 | 70 | 1131 | FERTIL STERIL |
| RODGERS K E | 1998 | 12 | 2443 | HUM REPROD |
| RODGERS K E | 1996 | 9 | 388 | J INVEST SURG |
| RODGERS K E | 1997 | 10 | 31 | J INVEST SURG |
| SHEAR L | ! | | | |
| SITTER T | 1965 | 272 | 123 | NEW ENGL J MED |
| - | 1999 | 82 | 1171 | THROMB HAEMOSTASIS |
| STOUT A L | ! | 146 | 73 | AM J OBSTET GYNECOL |
| STRICKER B | ! | 178 | - | J AM COLL SURGEONS |
| THOMAS S | 1997 | 29 | 246 | AM J KIDNEY DIS |
| | | | | |

| THORNTON M H | 1998 | 13 | 1480 | HUM REPROD |
|---------------------|------|----|------|----------------------|
| TOPLEY N | 1994 | 14 | S28 | PERITON DIALYSIS INT |
| TRIMBOSKEMPER T C M | 1989 | 51 | 1053 | FERTIL STERIL |
| TULANDI T | 1998 | 92 | 766 | OBSTET GYNECOL |
| VERCO S J S | 1999 | 14 | 275 | HUM REPROD |
| WEINSTEIN W M | 1974 | 10 | 1250 | INFECT IMMUN |
| WISEMAN D M | 1998 | 70 | 702 | FERTIL STERIL |

- L3 ANSWER 28 OF 29 SCISEARCH COPYRIGHT 2002 ISI (R)
- AN 2000:610679 SCISEARCH
- GA The Genuine Article (R) Number: 341MW
- TI Use of fibrinolytic agents in the **prevention** of postoperative adhesion formation
- AU Hellebrekers B W J (Reprint); TrimbosKemper T C M; Trimbos J B M Z; Emeis J J; Kooistra T
- CS LEIDEN UNIV, MED CTR, DEPT GYNECOL, POB 9600, NL-2300 RC LEIDEN, NETHERLANDS (Reprint); TNO, GAUBIUS LAB, LEIDEN, NETHERLANDS
- CYA NETHERLANDS
- SO FERTILITY AND STERILITY, (AUG 2000) Vol. 74, No. 2, pp. 203-212. Publisher: ELSEVIER SCIENCE INC, 655 AVENUE OF THE AMERICAS, NEW YORK, NY 10010.
 - ISSN: 0015-0282.
- DT General Review; Journal
- FS LIFE; CLIN
- LA English

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- REC Reference Count: 103
 - Objective: To review the events leading to the formation of adhesions, to describe the development of fibrinolytic agents, to review more than a century of research on the use of fibrinolytic agents in adhesion prevention, and to look at future aspects of adhesion prevention.

Results: A better understanding of the pathogenesis of adhesion formation has resulted in the use of fibrinolytic agents in their prevention. Fibrinolytic agents promote fibrinolytic activity during the early period after peritoneal trauma during which an increased formation of fibrin is seen in combination with a deficiency of endogenous fibrinolytic activity. Initially, chemical attacks on fibrin (fibrolysin and hypertonic glucose), foreign digestive ferments (pepsin, trypsin, and papain), and stimulation of intraperitoneal leuikocytosis (amniotic fluid) were used. Development of new thrombolytic agents was soon followed by experiments in animal adhesion models and clinical studies to examine their antiadhesion properties. Plasmin preparations (plasmin, actase, and fibrinolysin) and plasmin activators (streptokinase, urokinase, and tissue-type plasminogen activator) were found to be efficacious in preventing adhesion formation in the: greater part of reviewed animal and clinical studies.

Conclusion(s): From the current literature, it can be concluded that postoperative intraperitoneal administration of thrombolytic agents can significantly decrease adhesion formation. Given the large number of experimental studies in animals, future studies should focus on the clinical use of fibrinolytic agents in the **prevention** of postsurgical adhesion formation. (C) 2000 by American Society for Reproductive Medicine.

- CC OBSTETRICS & GYNECOLOGY; REPRODUCTIVE BIOLOGY
- ST Author Keywords: adhesion **prevention**; fibrinolytic agents; fibrolysin; pepsin; trypsin; papain; streptokinase; plasmin; urokinase; tissue plasminogen activator
- STP KeyWords Plus (R): TISSUE-PLASMINOGEN-ACTIVATOR; ACUTE
 MYOCARDIAL-INFARCTION; CHRONIC PELVIC PAIN; INTRAPERITONEAL
 ADHESIONS; INTRAABDOMINAL ADHESIONS; POSTSURGICAL
 ADHESIONS; MESOTHELIAL CELLS; ELECTIVE SURGERY; TUBAL SURGERY;
 RABBIT MODEL

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| MENZIES D | 1991 | 172 | 362 | SURG GYNECOL OBSTET |
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| MONTZ F J | 1991 | 165 | 1539 | AM J OBSTET GYNECOL |
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| RAFTERY A T | 1981 | 13 | 397 | EUR SURG RES |
| REA C E | 1933 | 31 | 1060 | P SOC EXP BIOL MED |
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| SCOTTCOOMBES D | 1995 | 82 | 414 | BRIT J SURG |
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| SIEVERS S | 1981 | 99 | 27 | FORTSCHR MED |
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| STEEGE J F | 1991 | 165 | 278 | AM J OBSTET GYNECOL |
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| TREUTNER K H | 1989 | 374 | 99 | LANGENBECK ARCH CHIR |
| TRIMBOSKEMPER T C M | 1985 | 43 | 395 | FERTIL STERIL |
| TRUSLER H M | 1931 | 22 | 983 | ARCH SURG-CHICAGO |
| TUCHMANN A | 1990 | 2 | 1041 | LANGENBECKS ARCH C S |
| VANGOOR H | 1996 | 28 | 287 | EUR SURG RES |
| VERREET P R | 1989 | 21 | 267 | EUR SURG RES |
| VERSTRAETE M | 1995 | 74 | 25 | THROMB HAEMOSTASIS |
| VIPOND M N | 1994 | 76 | 412 | ANN ROY COLL SURG |
| VIPOND M N | 1994 | 160 | 471 | EUR J SURG |
| VIPOND M N | 1990 | 335 | 1120 | LANCET |
| VONBENZER H | 1963 | 75 | 881 | WIEN KLIN WOCHENSCHR |
| WALTON R P | 1930 | 40 | 403 | J PHARMACOL EXP THER |
| WARREN S | 1928 | 6 | 860 | ARCH SURG-CHICAGO |
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| WHITTING H W | 1966 | 341 | 155 | VIRCHOWS ARCH PATHOL |
| WRIGHT L T | 1950 | 75 | 602 | P SOC EXP BIOL MED |
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